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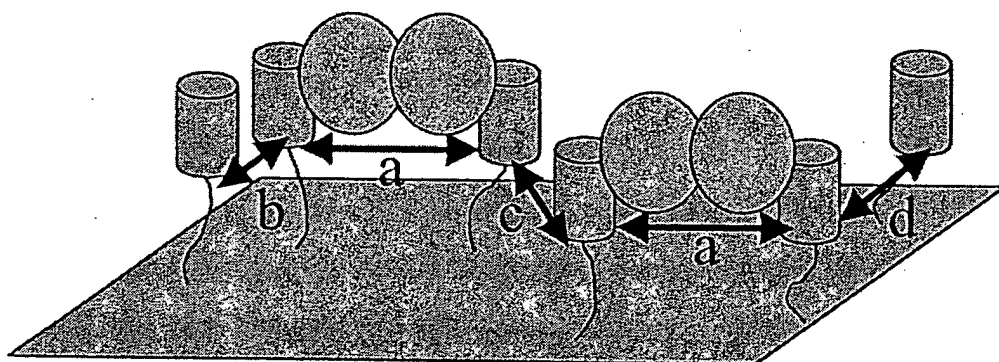
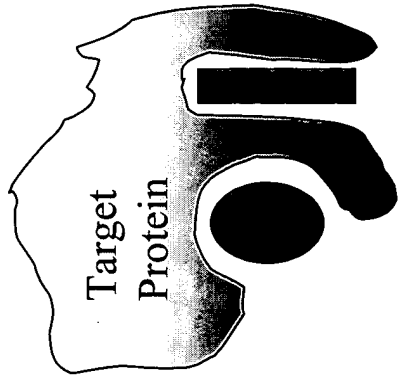
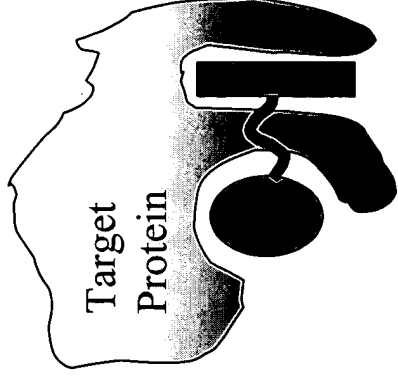


FIG. 1



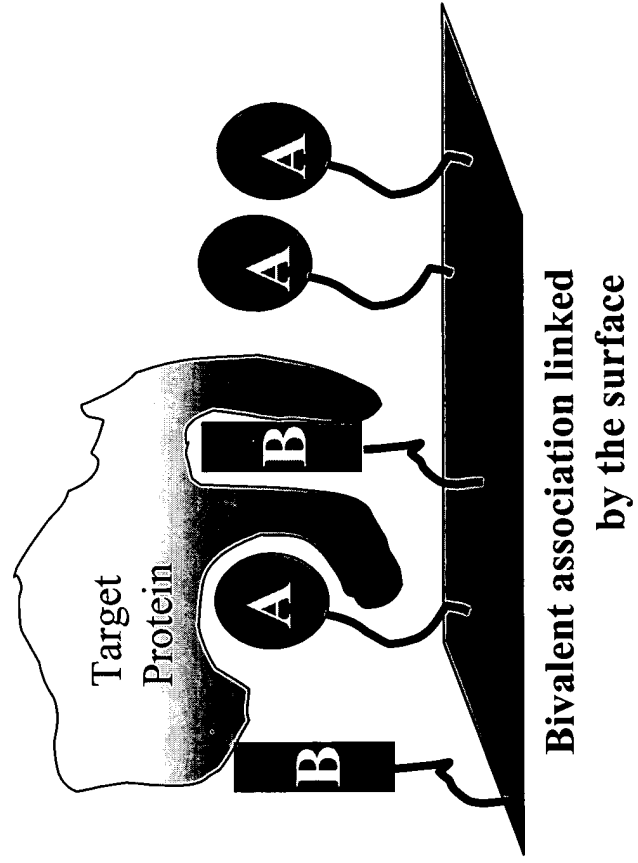
**Two low affinity  
monovalent ligands**

Fesik, Ellman, others



**High affinity  
bivalent ligand**

**FIG. 2**



**FIG. 3**

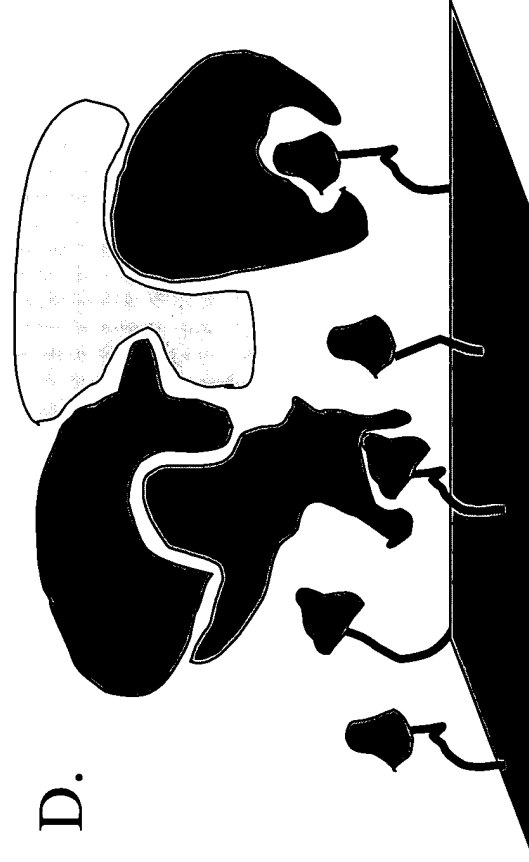
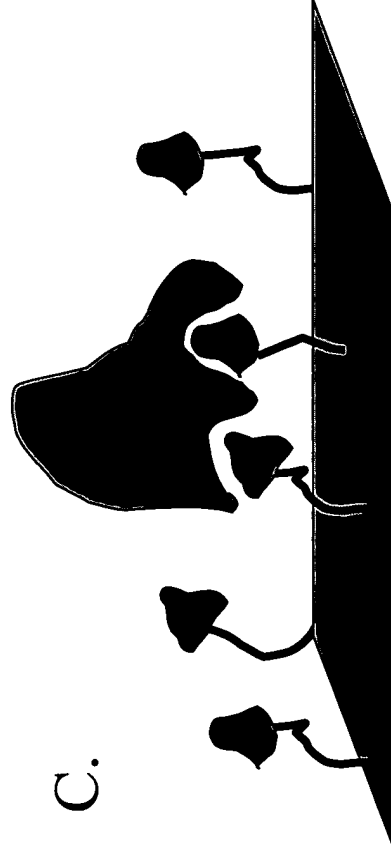
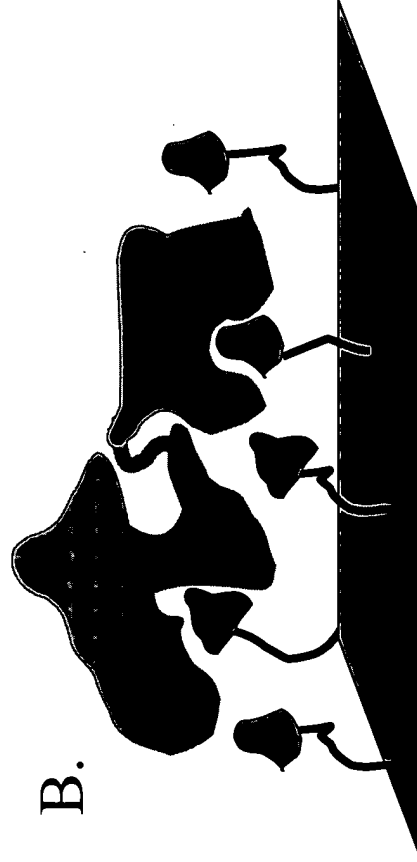
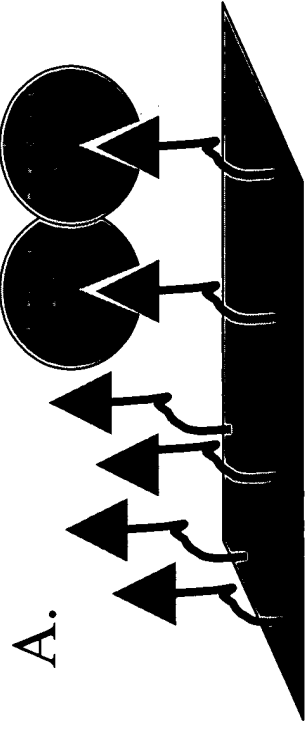
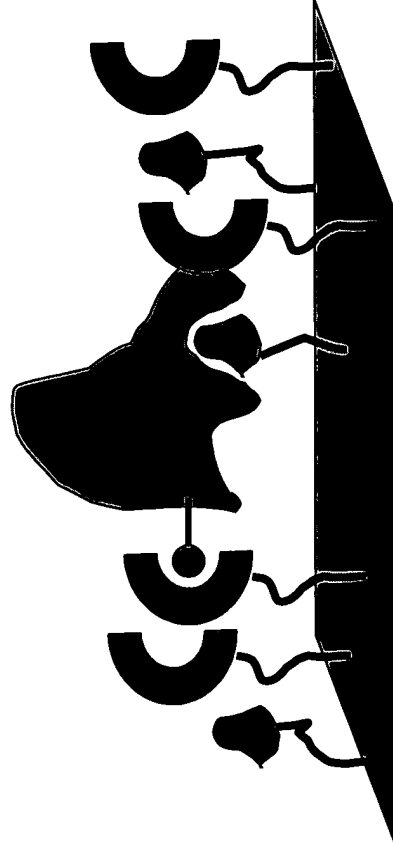
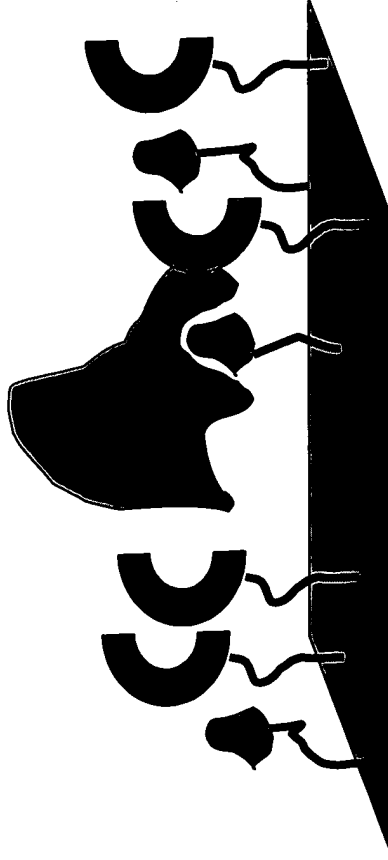


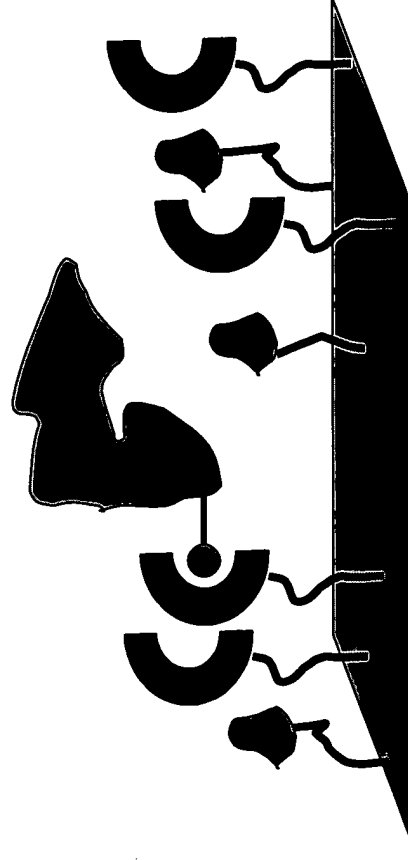
FIG. 4



High affinity binding of a particular post-translationally modified protein through a two-point contact.



The same protein lacking the modification would be bound weakly via a one-point contact.



Other proteins containing the same modification would be bound weakly via a one-point contact

FIG. 5

G80BP-A: YDQDMQNNTFDDDLFWKEGHR  
 G80BP-B: NEDWERDDQNPWDKLWMNRA  
 GAL4 AD: MDQTAYNAFGITTGMENTTMDDDVYNYLFDDDEDT

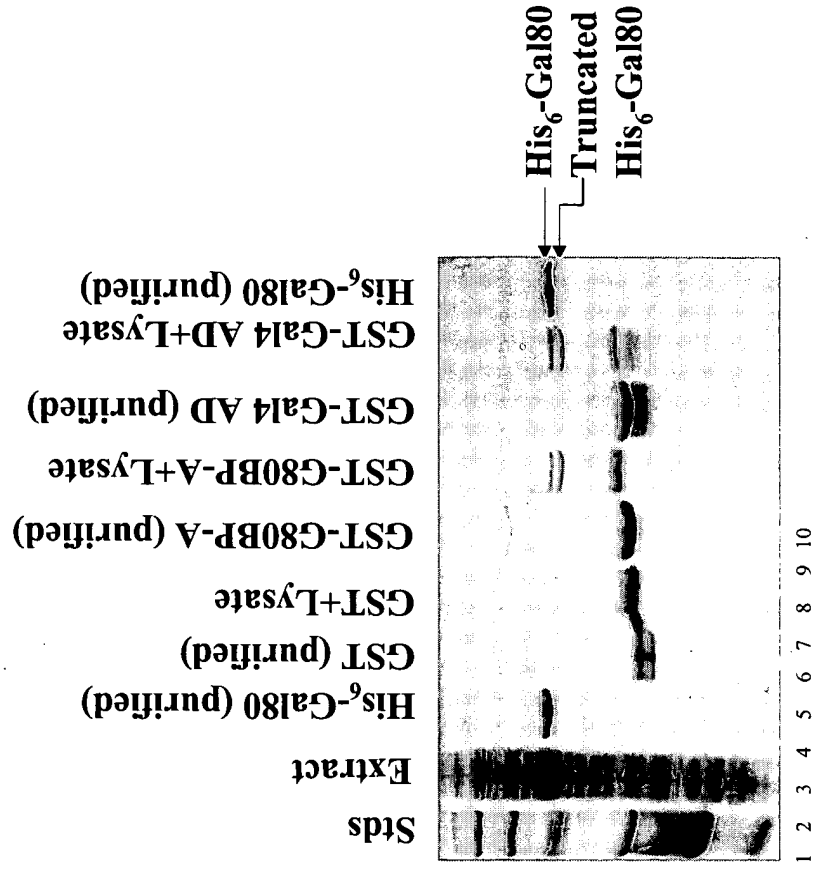
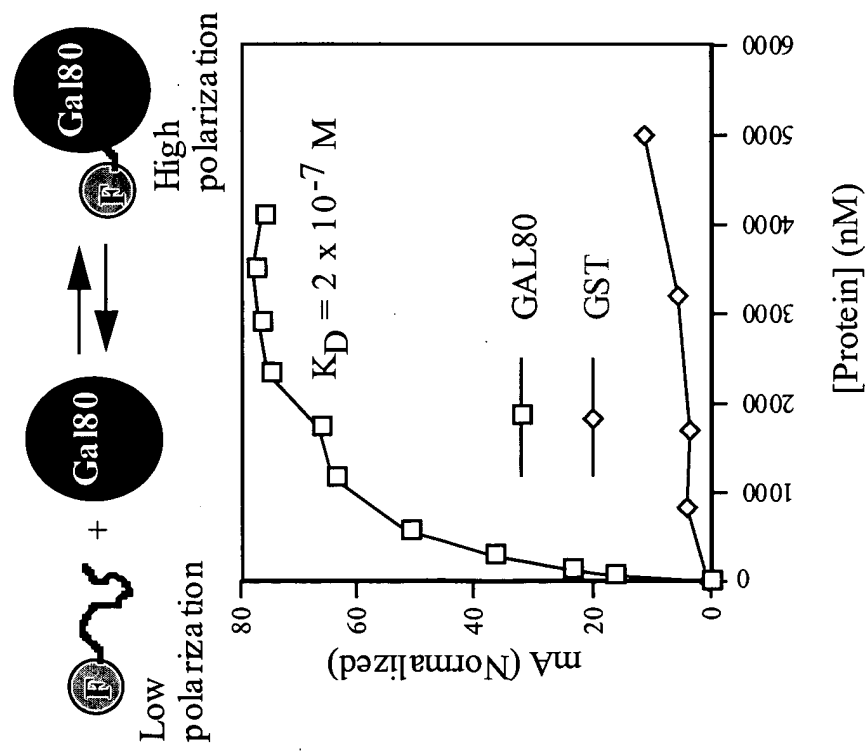
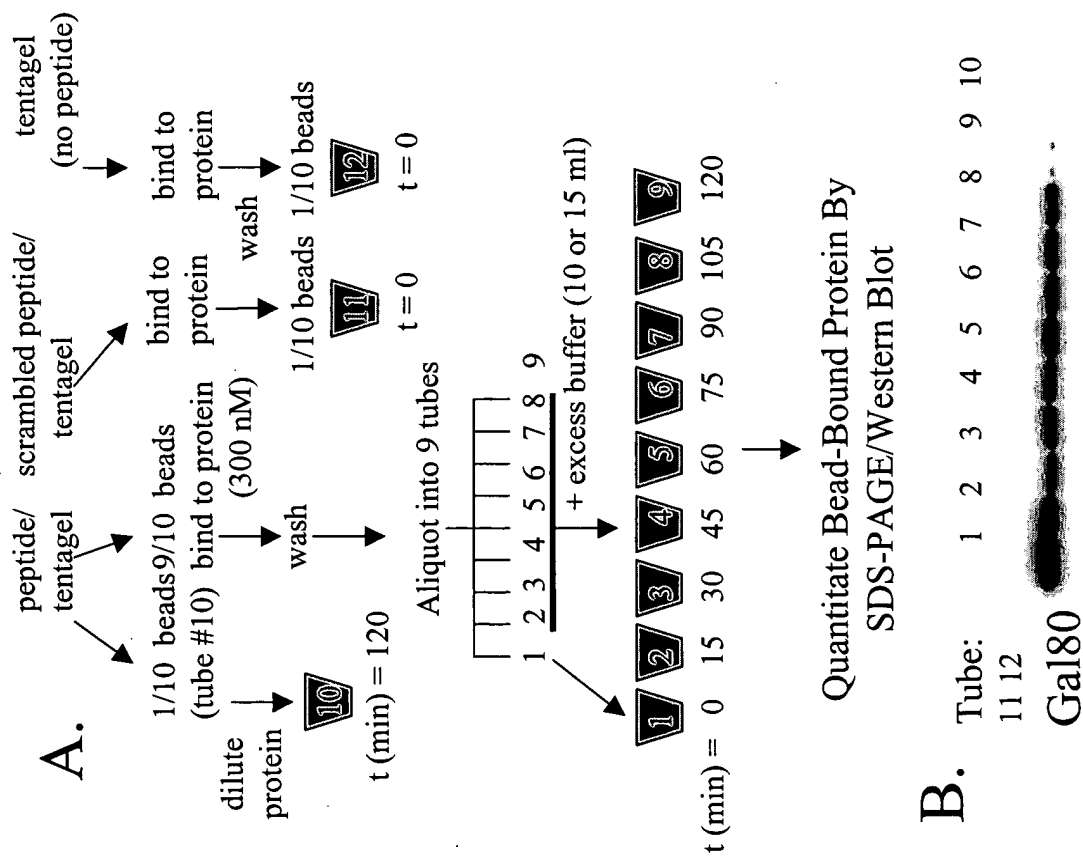


FIG. 6



**FIG. 7**



A. [His6Gal80] (nM): 8 4 2 1.5 1 0.75 0.4 0.3 0.2  
 His6Gal80 (ng): 200 100 50 37.5 25 17.5 10 7.5 5

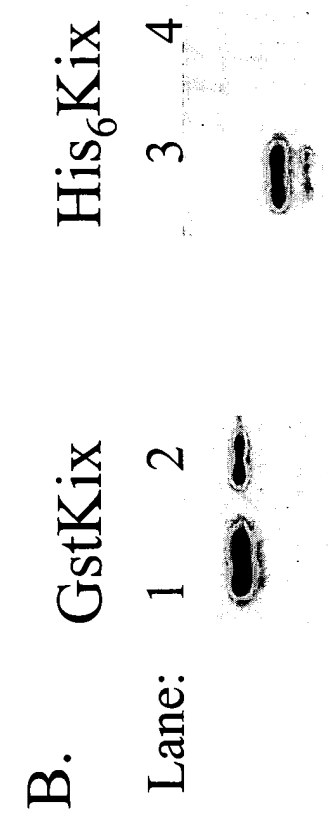
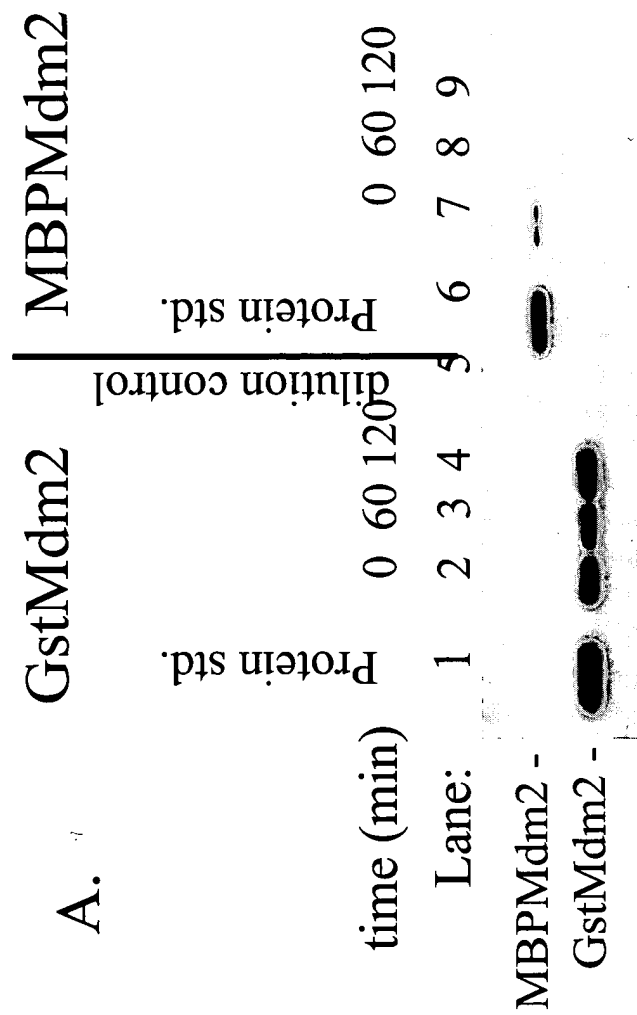


B.

His6Gal80 (ng) 84 63 42 30 21 15 10 5



FIG. 8



**FIG. 9**

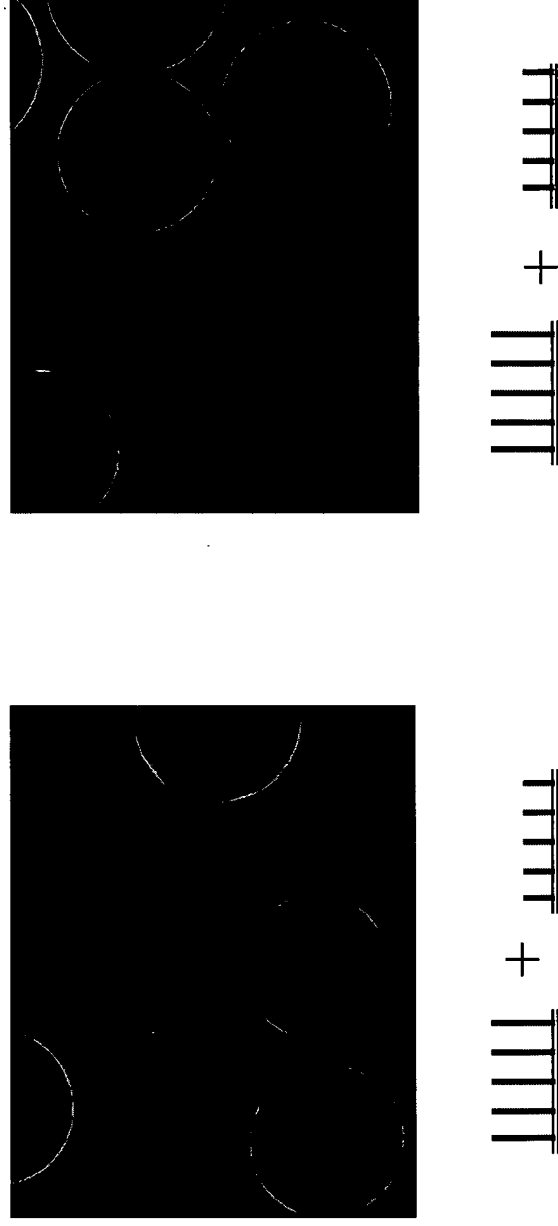
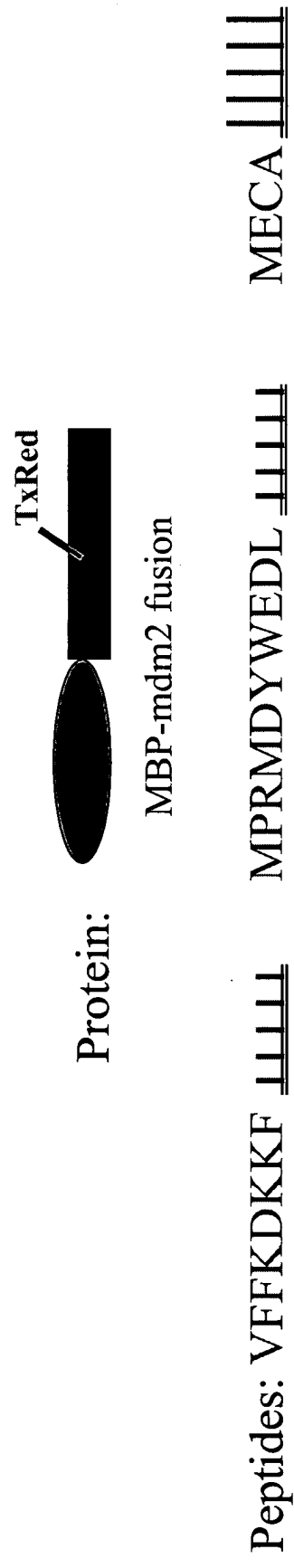
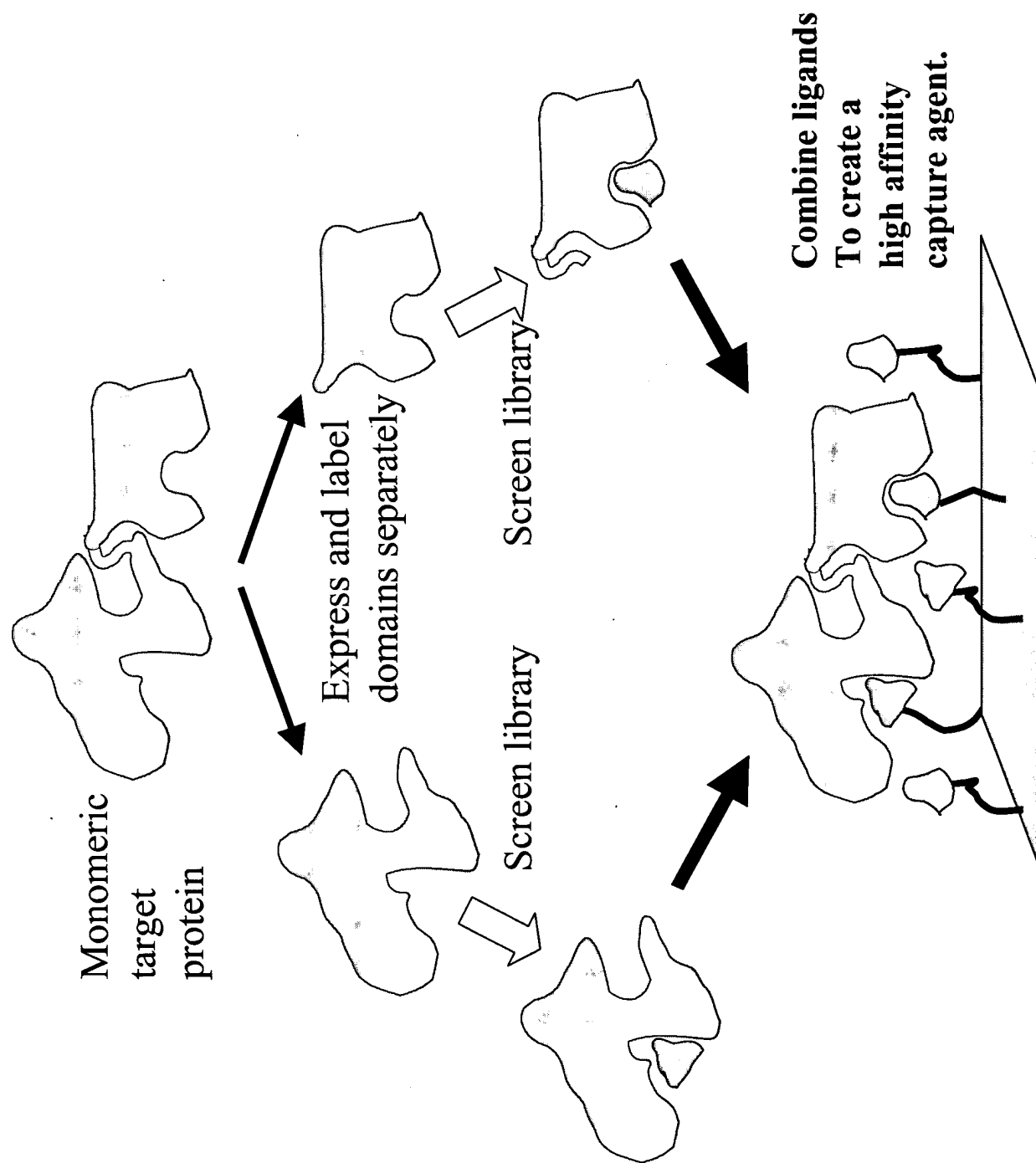
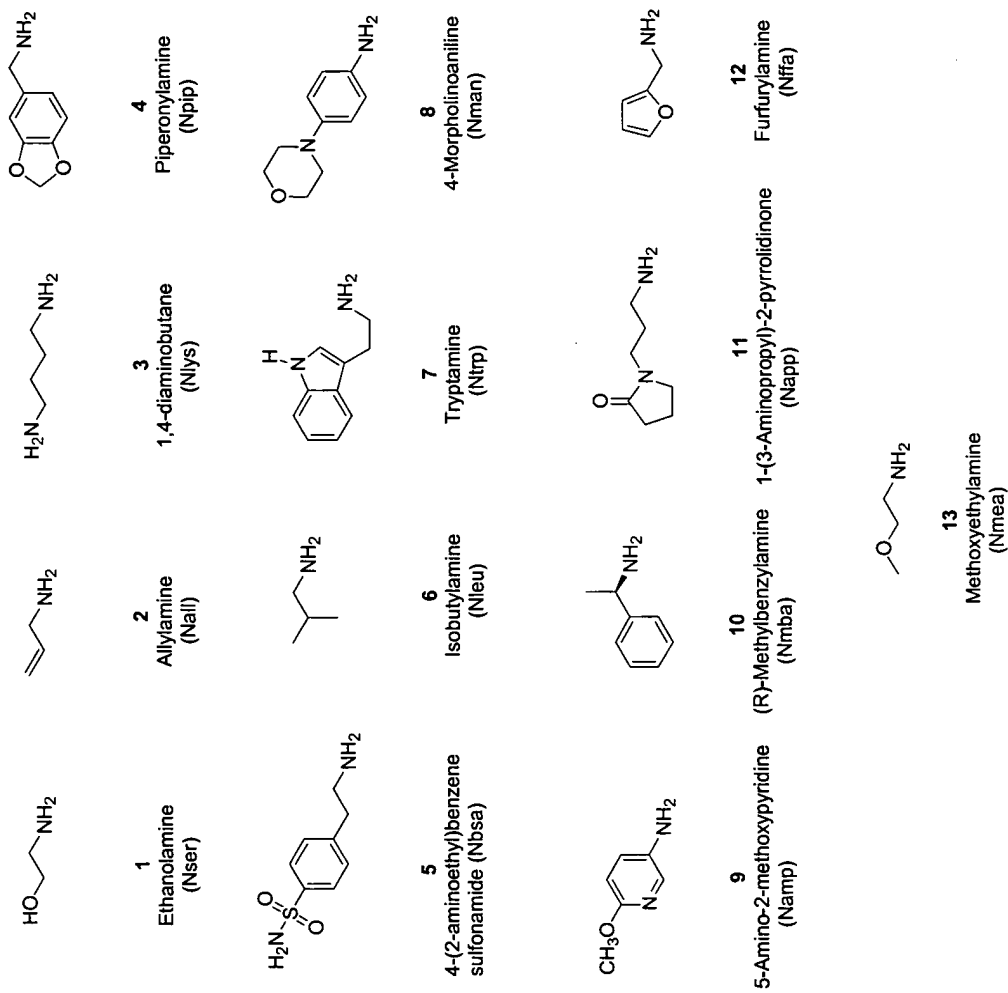


FIG. 10



**FIG. 11**



**FIG. 12**

- (1) Npip-Npip-Nbsa-Nser-Nliys-Npip-Nbsa-Nliys
- (2) Nbsa-Npip-Nliys-Nser-Nbsa-Nser-Nbsa-Npip
- (3) Nbsa-Nliys-Npip-Nser-Nbsa-Nser-Nall-Nliys
- (4) Nbsa-Nbsa-Nall-Nser-Nall-Npip-Nall-Npip
- (5) Nall-Nall-Nbsa-Nser-Nliys-Nbsa-Nser-Nall
- (6) Nliys-Nliys-Nser-Nser-Npip-Nall-Nbsa-Nser
- (7) Nliys-Nliys-Nser-Nser-Npip-Nbsa-Nall-Nbsa
- (8) Nall-Nliys-Nser-Nser-Nbsa-Nser-Nliys-Nliys
- (9) Npip-Nliys-Nall-Nser-Nbsa-Nser-Nbsa-Nall
- (10) Nbsa-Nliys-Npip-Nser-Nbsa-Nser-Nall-Nliys

**FIG. 13**

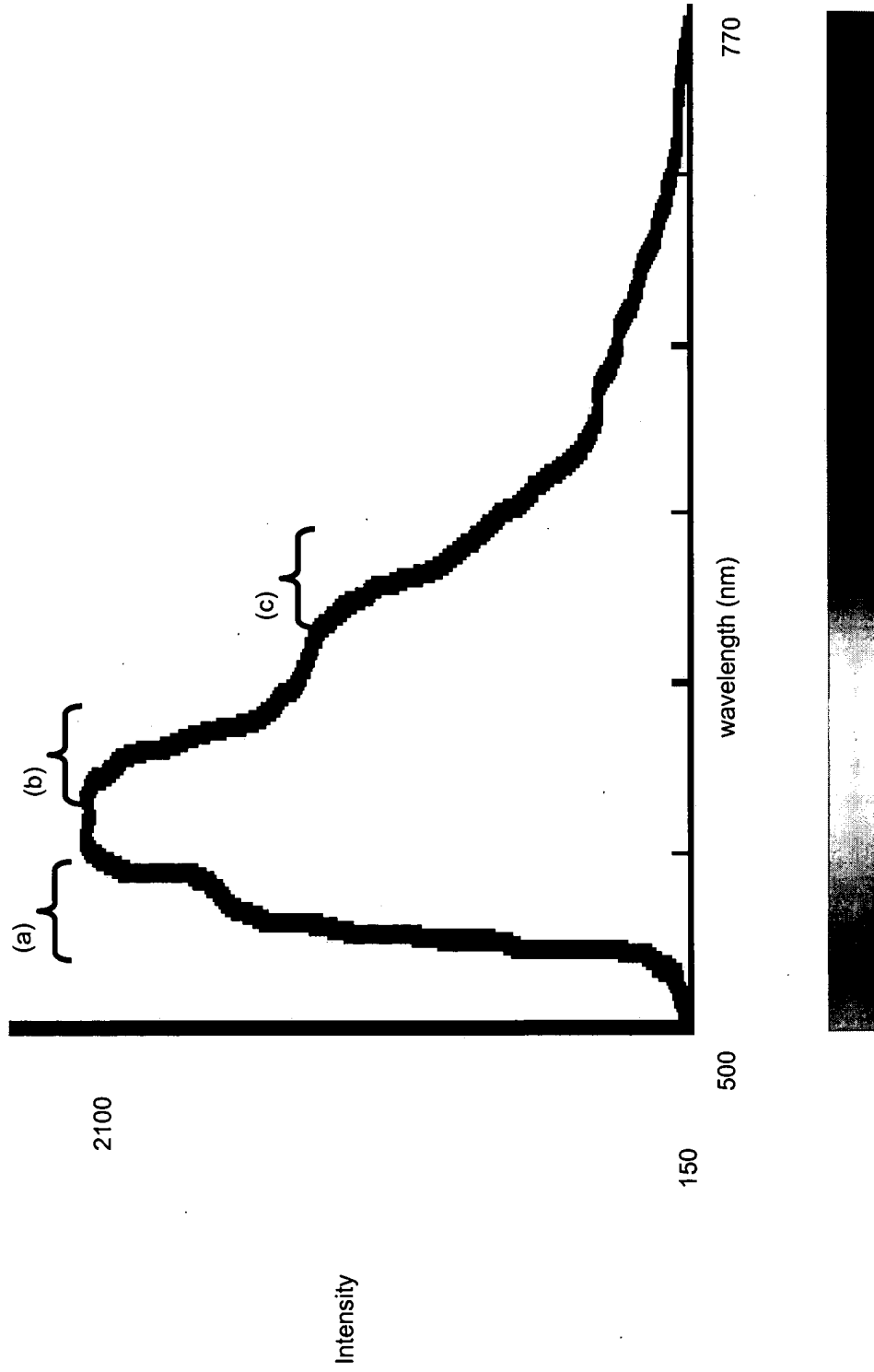


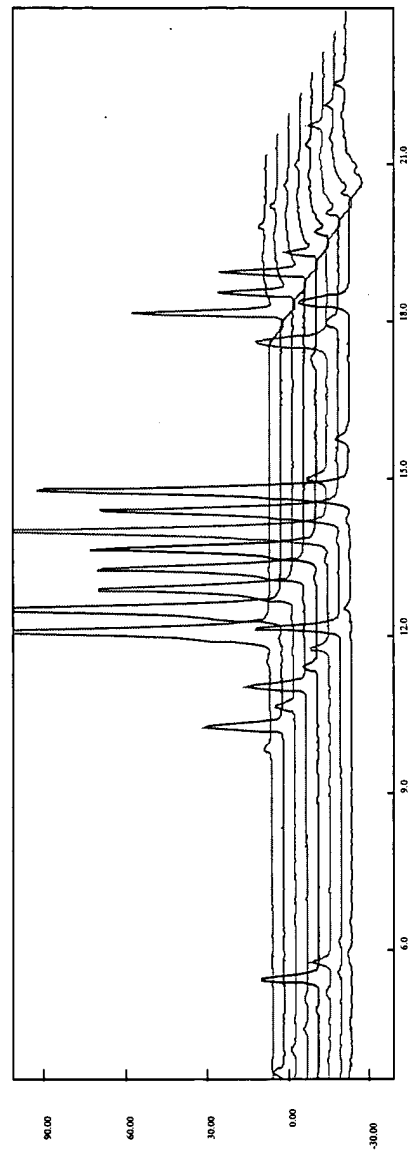
FIG. 14

(A)



Hit

(B)



(C)

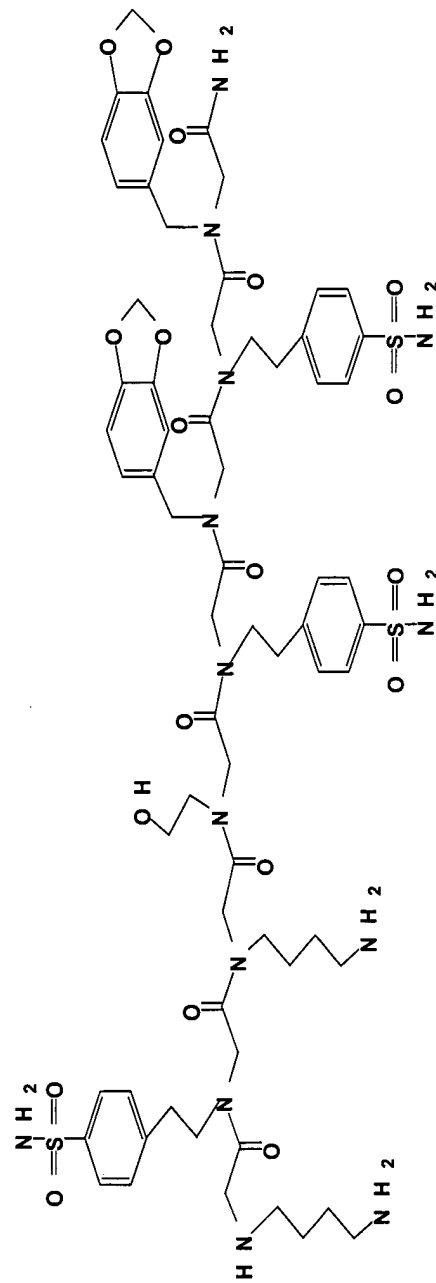


FIG. 15



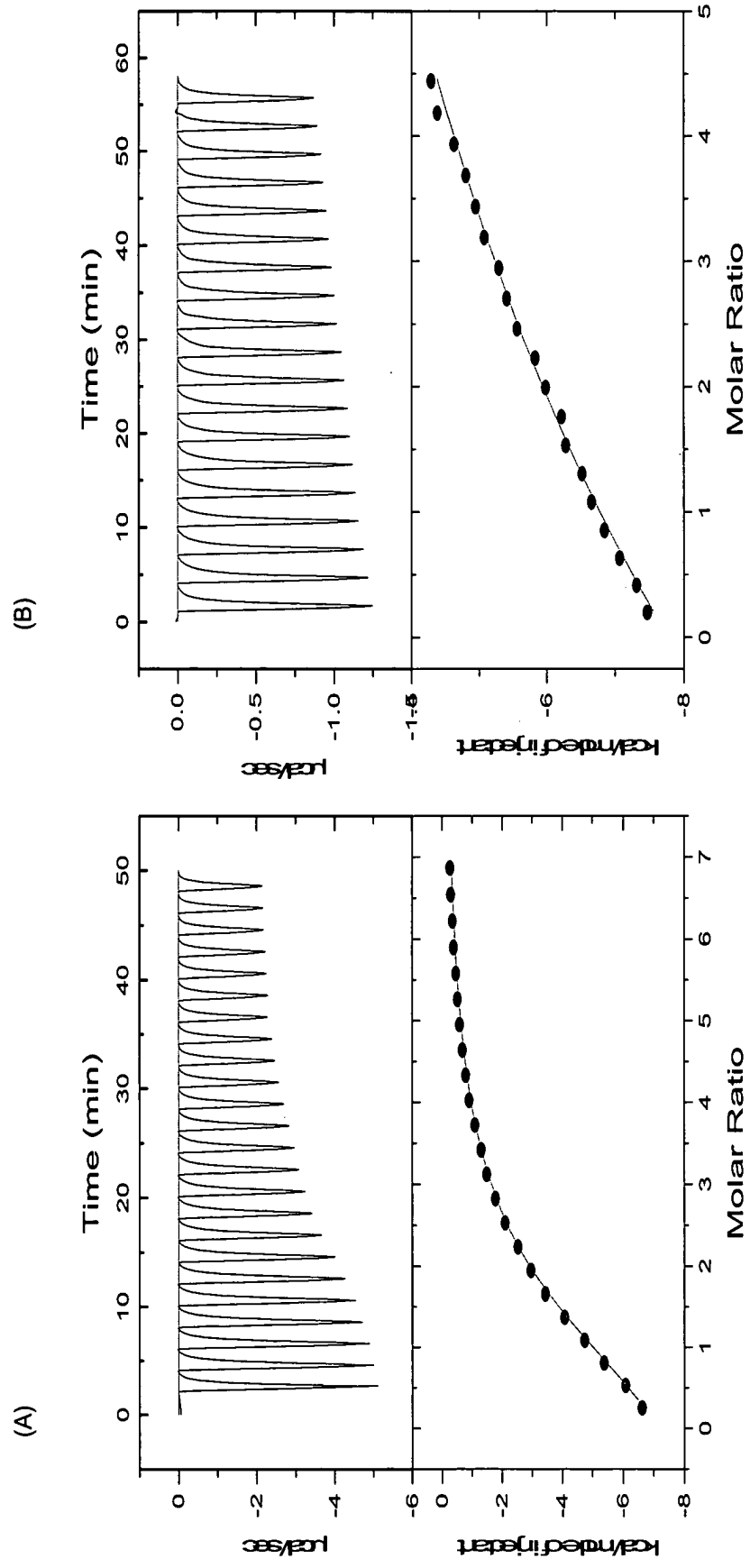


FIG. 16

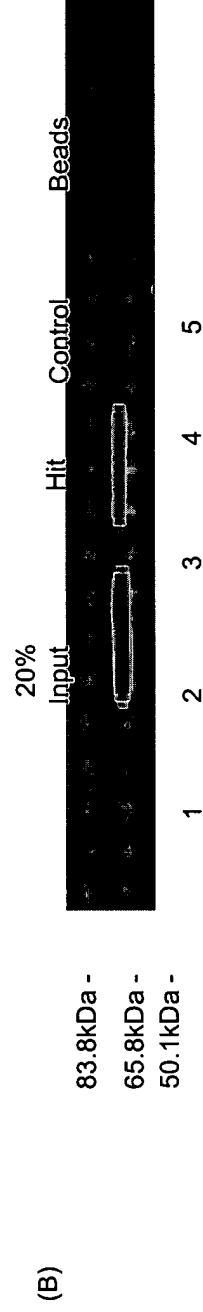
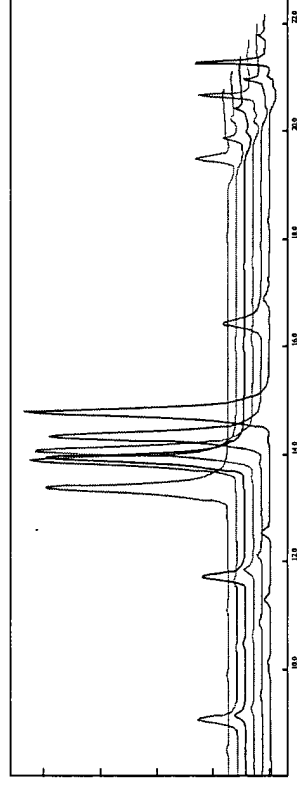


FIG. 17

(A)

- Nleu-Nbsa-Nlys-Napp-Ntrp-Nleu
- Nmba-Nleu-Napp-Nmba-Nbsa-Nffa
- Nlys-Nmea-Ntrp-Napp-Nleu-Nleu
- Nmea-Npip-Nmba-Nffa-Napp-Nleu
- Nleu-Nlys-Nmea-Nleu-Ntrp-Nmea
- Ntrp-Nffa-Ntrp-Nbsa-Napp-Npip
- Nffa-Nleu-Nmea-Npip-Nlys-Nleu
- Npip-Nmea-Nmba-Napp-Nleu-Napp
- Napp-Nmea-Nmba-Npip-Nffa-Nffa
- Nleu-Npip-Nbsa-Nmea-Nmba-Nmea

(B)



(C)

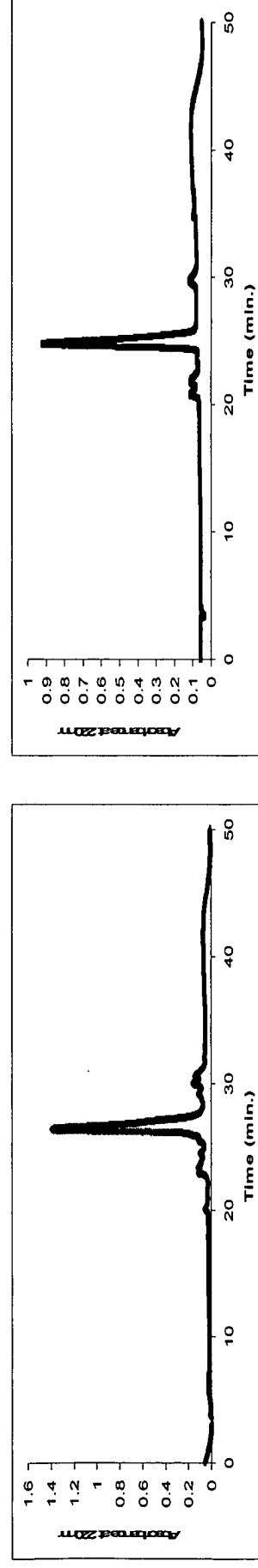


FIG. 18

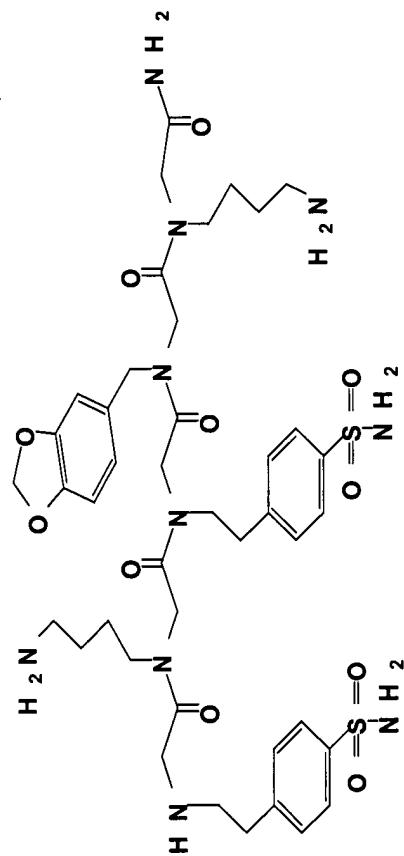
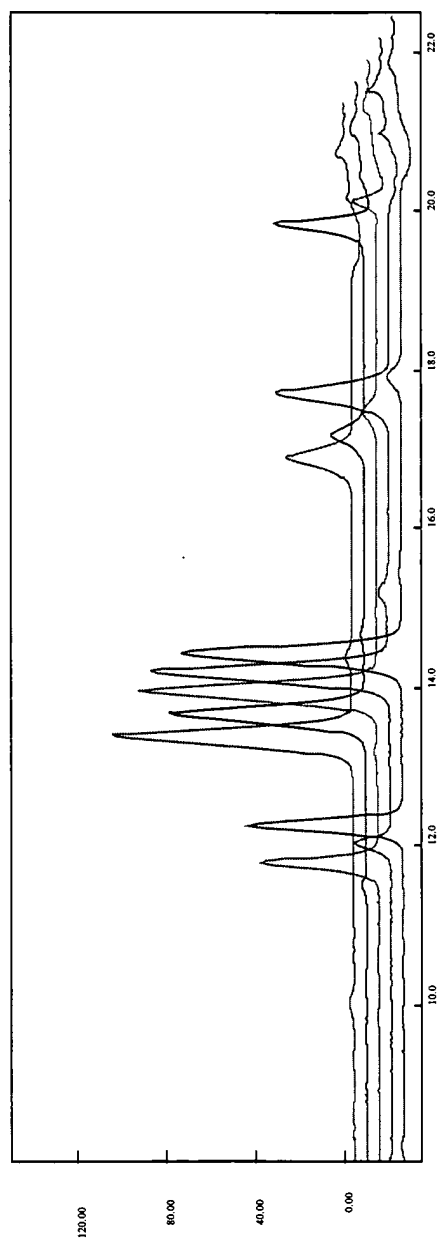
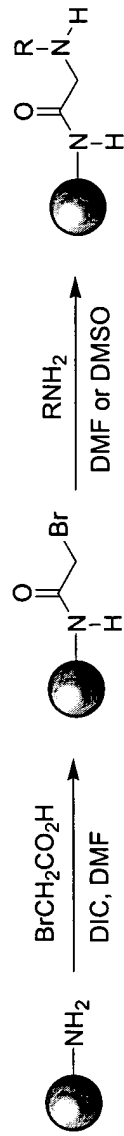


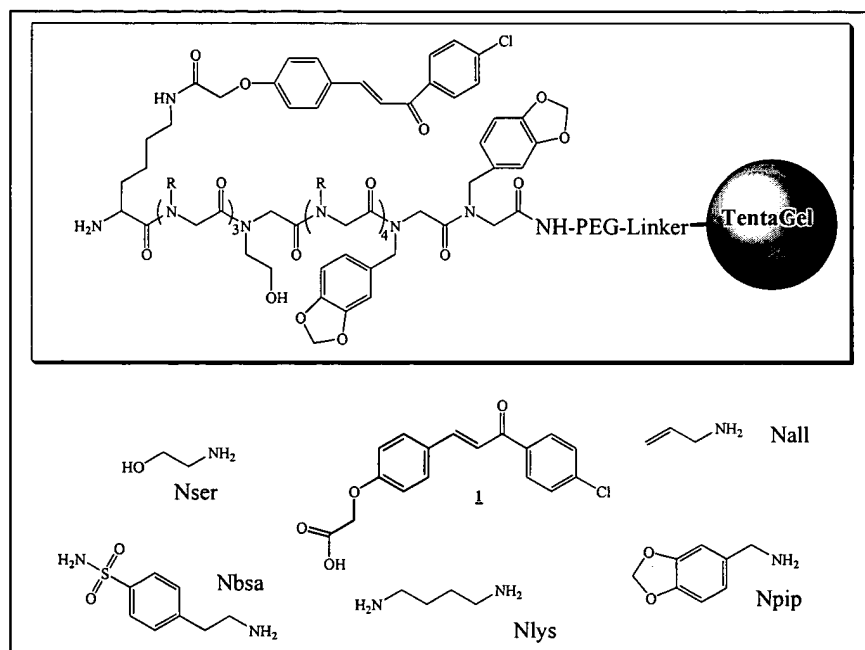
FIG. 19



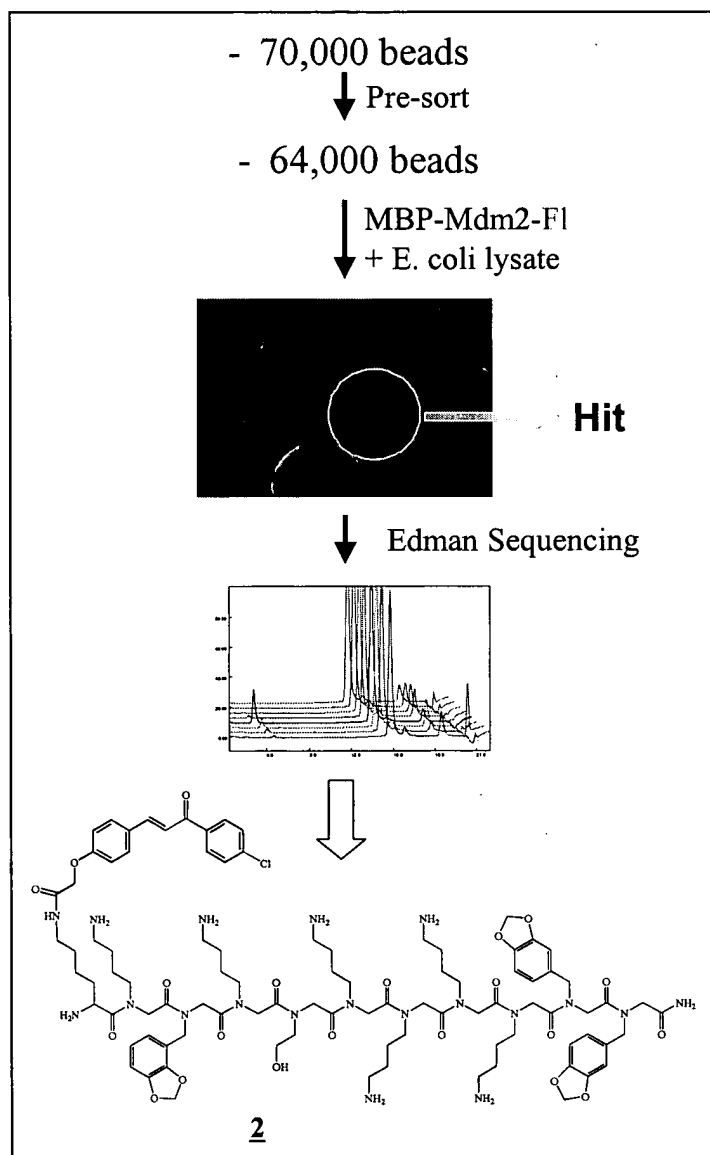


DIC :  $(^i\text{Pr})\text{N}=\text{C}=\text{N}(^i\text{Pr})$   
 R : any alkyl group

FIG. 21

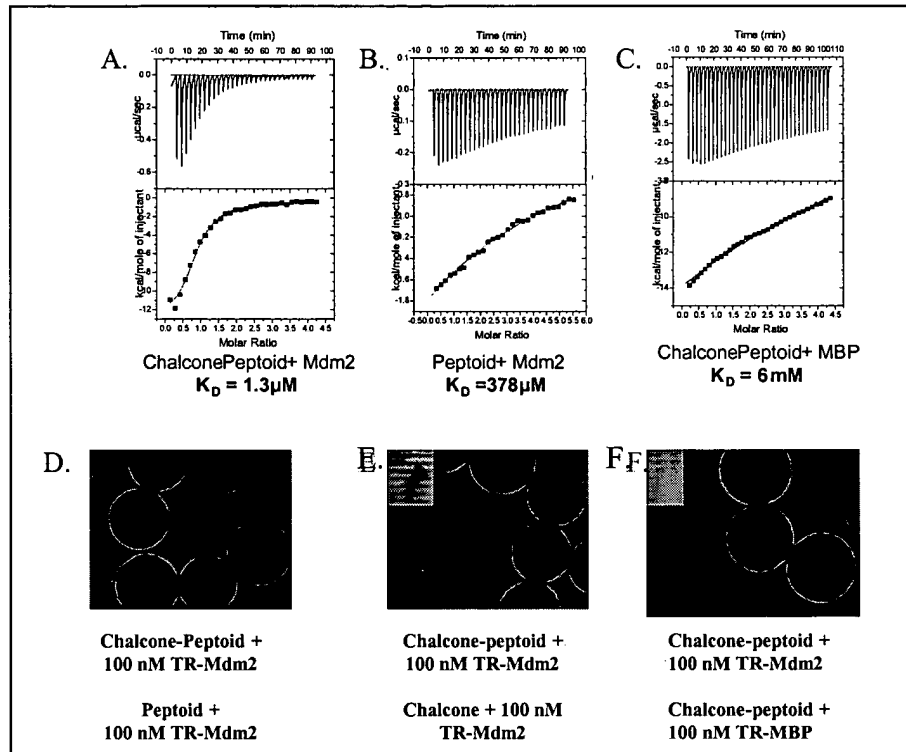


**FIG. 22**

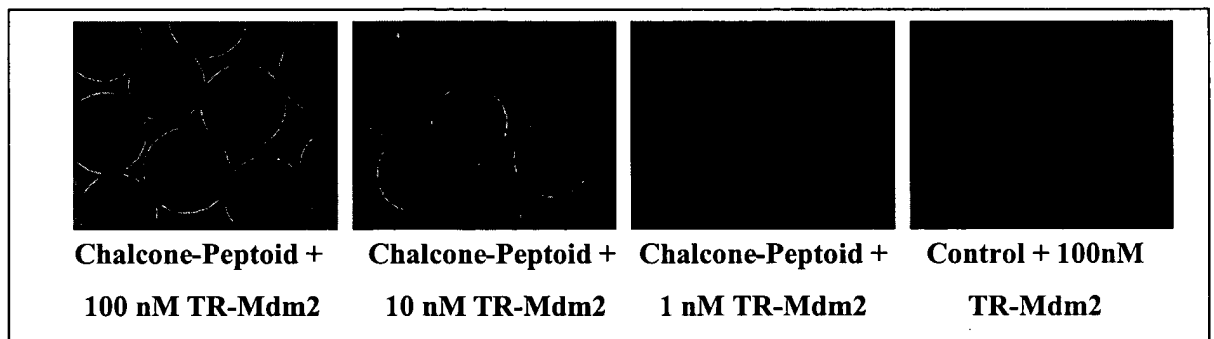


**FIG. 23A-23C**





**FIG. 24A-24F**



**FIG 25**

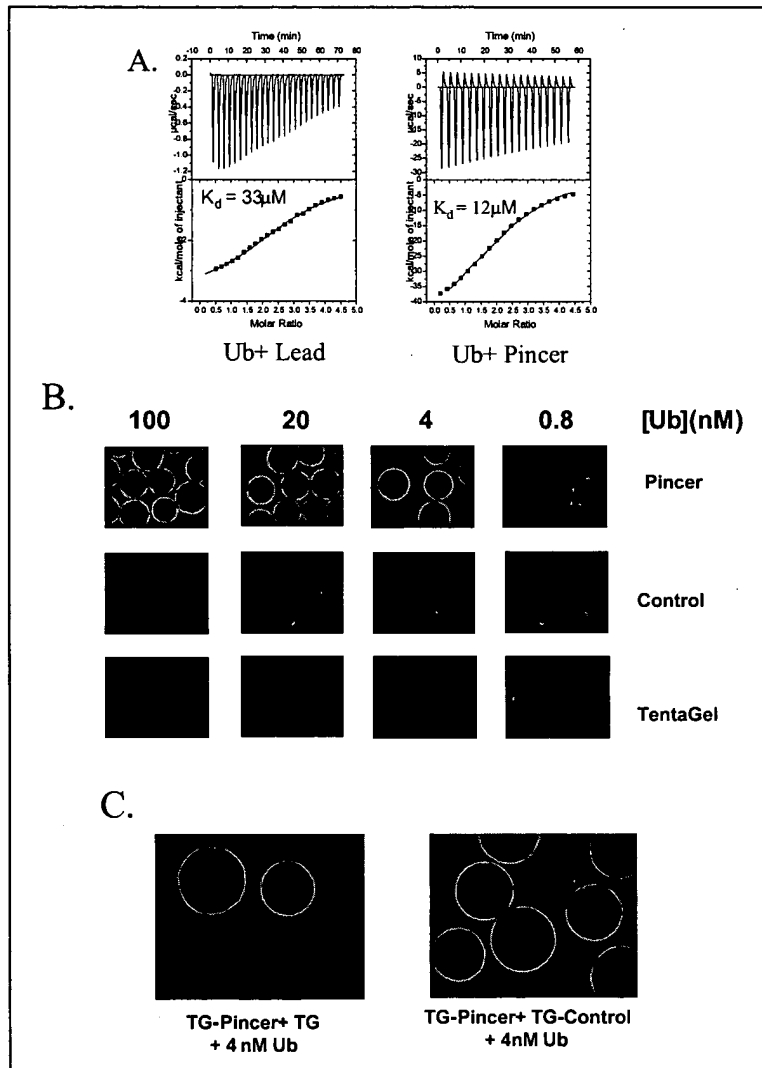


FIG. 26A-26C